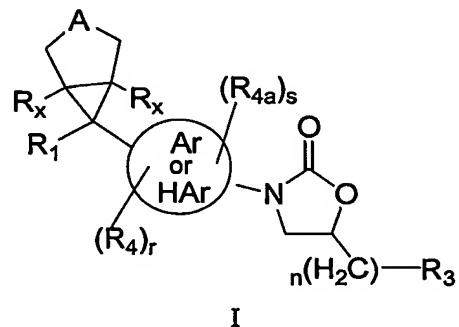


WHAT IS CLAIMED IS:

1. The present invention relates to compounds of formula I:

5



its enantiomer, diastereomer, or pharmaceutically acceptable salt, hydrate or prodrug thereof
10 wherein:

R₁ represents

- vi) hydrogen,
- vii) NR₅R₆,
- 15 viii) CR₇R₈R₉, C(R)₂OR₁₄, CH₂NHR₁₄,
- ix) C(=O)R₁₃, C(=NOH)H, C(=NOR₁₃)H, C(=NOR₁₃)R₁₃, C(=NOH)R₁₃, C(=O)N(R₁₃)₂,
C(=NOH)N(R₁₃)₂, NHC(=X₁)N(R₁₃)₂, (C=NH)R₇, N(R₁₃)C(=X₁)N(R₁₃)₂, COOR₁₃,
SO₂R₁₄, N(R₁₃)SO₂R₁₄, N(R₁₃)COR₁₄,
- x) (C₁₋₆alkyl)CN, CN, CH=C(R)₂, (CH₂)_pOH, C(=O)CHR₁₃, C(=NR₁₃)R₁₃,
20 NR₁₀C(=X₁)R₁₃; or
- vi) C₅₋₁₀ heterocycle optionally substituted with 1-3 groups of R₇, which may be attached
through either a carbon or a heteroatom;
- 25 A represents NR, O, or S(O)p;

Ar
or
HAr

represents aryl or heteroaryl, heterocycle, heterocyclyl or heterocyclic, provided that in the case of a heteroaryl, heterocycle, heterocyclyl or heterocyclic, the cyclopropyl is not attached to a nitrogen atom on the ring;

5 R_x represents hydrogen or C₁₋₆ alkyl;

R₃ represent

- i) NR₁₃(C=X₂)R₁₂,
- ii) NR₁₃(C=X₁)R₁₂,
- 10 iii) NR₁₃SO₂R₁₄,
- iv) N(R₁₃)heteroaryl,
- v) NR₁₃(CHR₁₃)₀₋₄aryl,
- vi) NR₁₃(CHR₁₃)₀₋₄heteroaryl,
- vii) S(CHR₁₃)₀₋₄aryl,
- 15 viii) S(CHR₁₃)₀₋₄heteroaryl,
- ix) O(CHR₁₃)₀₋₄aryl,
- x) O(CHR₁₃)₀₋₄heteroaryl,
- xi) NOH(C=X₁)R₁₂,
- xii) -OC=N(OCOaryl) C₁₋₆ alkyl
- 20 xiii) -OC=N(OH) C₁₋₆ alkyl
- xiv) C₅₋₁₀ heteroaryl which may be attached through either a carbon or a heteroatom; said aryl and heteroaryl optionally substituted with 1-3 groups of R₇,

R₄, and R_{4a}, independently represent

- 25 v) hydrogen,
- vi) halogen,
- vii) C₁₋₆ alkoxy, or
- viii) C₁₋₆ alkyl
- 30 r and s independently are 1-3, with the provision that when (R_{4a})_s and (R₄)_r are attached to an Ar or HAr ring the sum of r and s is less than or equal to 4;

R₅ and R₆ independently represent

- xiii) hydrogen,
- xiv) C₁₋₆ alkyl optionally substituted with 1-3 groups of halogen, CN, OH, C₁₋₆ alkoxy, amino, imino, hydroxyamino, alkoxyamino, C₁₋₆ acyloxy, C₁₋₆ alkylsulfenyl, C₁₋₆ alkylsulfinyl, C₁₋₆ alkylsulfonyl, aminosulfonyl, C₁₋₆ alkylaminosulfonyl, C₁₋₆ dialkylaminosulfonyl, 4-morpholinylsulfonyl, phenyl, pyridine, 5-isoxazolyl, ethylenyloxy, or ethynyl, said phenyl and pyridine optionally substituted with 1-3 halogen, CN, OH, CF₃, C₁₋₆ alkyl or C₁₋₆ alkoxy;
- xv) C₁₋₆ acyl optionally substituted with 1-3 groups of halogen, OH, SH, C₁₋₆ alkoxy, naphthalenoxy, phenoxy, amino, C₁₋₆ acylamino, hydroxylamino, alkoxyamino, C₁₋₆ acyloxy, aralkyloxy, phenyl, pyridine, C₁₋₆ alkylcarbonyl, C₁₋₆ alkylamino, C₁₋₆ dialkylamino, C₁₋₆ hydroxyacyloxy, C₁₋₆ alkylsulfenyl, phthalimido, maleimido, succinimido, said phenoxy, phenyl and pyridine optionally substituted with 1-3 groups of halo, OH, CN, C₁₋₆ alkoxy, amino, C₁₋₆ acylamino, CF₃ or C₁₋₆ alkyl;
- xvi) C₁₋₆ alkylsulfonyl optionally substituted with 1-3 groups of halogen, OH, C₁₋₆ alkoxy, amino, hydroxylamino, alkoxyamino, C₁₋₆ acyloxy, or phenyl; said phenyl optionally substituted with 1-3 groups of halo, OH, C₁₋₆ alkoxy, amino, C₁₋₆ acylamino, CF₃ or C₁₋₆ alkyl;
- xvii) arylsulfonyl optionally substituted with 1-3 of halogen, C₁₋₆ alkoxy, OH or C₁₋₆ alkyl;
- xviii) C₁₋₆ alkoxycarbonyl optionally substituted with 1-3 of halogen, OH, C₁₋₆ alkoxy, C₁₋₆ acyloxy, or phenyl, said phenyl optionally substituted with 1-3 groups of halo, OH, C₁₋₆ alkoxy, amino, C₁₋₆ acylamino, CF₃ or C₁₋₆ alkyl;
- xix) aminocarbonyl, C₁₋₆ alkylaminocarbonyl or C₁₋₆ dialkylaminocarbonyl, said alkyl groups optionally substituted with 1-3 groups of halogen, OH, C₁₋₆ alkoxy or phenyl
- xx) five to six membered heterocycles optionally substituted with 1-3 groups of halogen, OH, CN, amino, C₁₋₆ acylamino, C₁₋₆ alkylsulfonylamino, C₁₋₆ alkoxycarbonylamino, C₁₋₆ alkoxy, C₁₋₆ acyloxy or C₁₋₆ alkyl, said alkyl optionally substituted with 1-3 groups of halogen, or C₁₋₆ alkoxy;
- xxi) C₃₋₆ cycloalkylcarbonyl optionally substituted with 1-3 groups of halogen, OH, C₁₋₆ alkoxy or CN;
- xxii) benzoyl optionally substituted with 1-3 groups of halogen, OH, C₁₋₆ alkoxy, C₁₋₆ alkyl, CF₃, C₁₋₆ alkanoyl, amino or C₁₋₆ acylamino;
- xxiii) pyrrolylcarbonyl optionally substituted with 1-3 of C₁₋₆ alkyl;

xxiv) C₁₋₂ acyloxyacetyl where the acyl is optionally substituted with amino, C₁₋₆ alkylamino, C₁₋₆ dialkylamino, 4-morpholino, 4-aminophenyl, 4-(dialkylamino)phenyl, 4-(glycylamino)phenyl; or
 R₅ and R₆ taken together with any intervening atoms can form a 3 to 7 membered
 5 heterocyclic ring containing carbon atoms and 1-2 heteroatoms independently chosen from O, S, SO, SO₂, N, or NR₈;

R₇ represent
 iii) hydrogen, halogen, CN, CO₂R, CON(R)₂, CHO, CH₂NHAc, C(=NOR), OH, C₁₋₆ 10 alkoxy, C₁₋₆ alkyl, alkenyl, hydroxy C₁₋₆ alkyl, (CH₂)₁₋₃NHC(O)C₁₋₆ alkyl, (CH₂)₁₋₃N(C₁₋₆ alkyl)₂
 iv) (CH₂)_namino, (CH₂)_nC₁₋₆ alkylamino, C₁₋₆ dialkylamino, hydroxylamino or C₁₋₂ 15 alkoxyamino all of which can be optionally substituted on the nitrogen with C₁₋₆ acyl, C₁₋₆ alkylsulfonyl or C₁₋₆ alkoxy carbonyl, said acyl and alkylsulfonyl optionally substituted with 1-2 of halogen or OH;

R₈ and R₉ independently represents
 iv) H, CN,
 v) C₁₋₆ alkyl optionally substituted with 1-3 halogen, CN, OH, C₁₋₆ alkoxy, C₁₋₆ 20 acyloxy, or amino,
 vi) phenyl optionally substituted with 1-3 groups of halogen, OH, C₁₋₆ alkoxy; or
 R₇ and R₈ taken together can form a 3-7 membered carbon ring optionally interrupted with 1-2 heteroatoms chosen from O, S, SO, SO₂, NH, and NR₈;
 25 X₁ represents O, S or NR₁₃, NCN, NCO₂R₁₆, or NSO₂R₁₄
 X₂ represents O, S, NH or NSO₂R₁₄;
 30 R₁₀ represents hydrogen, C₁₋₆ alkyl or CO₂R₁₅;
 R₁₂ represents hydrogen, C₁₋₆ alkyl, NH₂, OR, CHF₂, CHCl₂, CR₂Cl, (CH₂)_nSR, (CH₂)_nCN, (CH₂)_nSO₂R, (CH₂)_nS(O)R, C₁₋₆ alkylamino, C₅₋₁₀ heteroaryl or C₁₋₆ dialkylamino, where

said alkyl may be substituted with 1-3 groups of halo, CN, OH or C₁₋₆ alkoxy, said heteroaryl optionally substituted with 1-3 groups of R₇;

Each R₁₃ represents independently hydrogen, C₁₋₆ alkyl, C₆₋₁₀ aryl, NR₅R₆, SR₈, S(O)R₈, S(O)₂R₈, CN, OH, C₁₋₆ alkylS(O)R, C₁₋₆ alkoxy carbonyl, hydroxycarbonyl, -OCOaryl, C₁₋₆ acyl, C₃₋₇ membered carbon ring optionally interrupted with 1-4 heteroatoms chosen from O, S, SO, SO₂, NH and NR₈ where said C₁₋₆ alkyl, aryl or C₁₋₆ acyl groups may be independently substituted with 0-3 halogens, hydroxy, N(R)₂, CO₂R, C₆₋₁₀ aryl, C₅₋₁₀ heteroaryl, or C₁₋₆ alkoxy groups;

When two R₁₃ groups are attached to the same atom or two adjacent atoms they may be taken together to form a 3-7 membered carbon ring optionally interrupted with 1-2 heteroatoms chosen from O, S, SO, SO₂, NH, and NR₈;

R represents hydrogen or C₁₋₆ alkyl;

R₁₄ represents amino, C₁₋₆ alkyl, C₁₋₆ haloalkyl, five to six membered heterocycles or phenyl, said phenyl and heterocycles optionally substituted with 1-3 group of halo, C₁₋₆ alkoxy, C₁₋₆ acylamino, or C₁₋₆ alkyl, hydroxy and/or amino, said amino and hydroxy optionally protected with an amino or hydroxy protecting group;

R₁₅ is C₁₋₆ alkyl or benzyl said benzyl optionally substituted with 1-3 groups of halo, OH, C₁₋₆ alkoxy, amino, C₁₋₆ acylamino, or C₁₋₆ alkyl;

R₁₆ is hydrogen, C₅₋₁₀ heteroaryl, C₆₋₁₀ aryl, said heteroaryl and aryl optionally substituted with 1-3 groups of R₇;

p represents 0-2 and

m, n, and q represents 0-1.

2. A compound according to claim 1 wherein R₁ represents H, NR₅R₆, CN, OH, C(R)₂OR₁₄, NHC(=X₁)N(R₁₃)₂, C(=NOH)N(R₁₃)₂, NR₁₀C(=X₁)R₁₃ or CR₇R₈R₉.

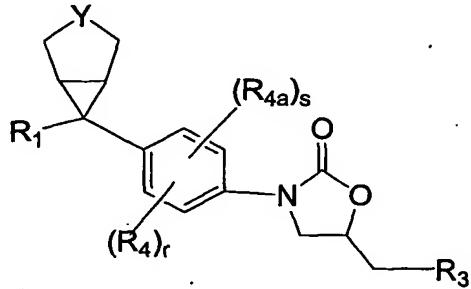
Ar
or
HAr

3. A compound according to claim 1 wherein
is phenyl, pyridine, pyrimidine, or piperidine.

4. A compound according to claim 3 wherein R₁ is NR₅R₆, or CN and
5 R₃ is NR₁₀C(=X₁)R₁₃, NR(C=X₁)R₁₂, C₅-10 heteroaryl, NH(CH₂)₀₋₄aryl, NH(CH₂)₀₋₄heteroaryl, said aryl and heteroaryl optionally substituted with 1-3 groups of R_a.

5. A compound according to claim 3 wherein R₃ is a C₅-10 heteroaryl
represented by  which represents an optionally substituted aromatic heterocyclic group
10 containing 1 to 4 nitrogen atoms and at least one double bond, and which is connected
through a bond on any nitrogen.

6. A compound according to claim 1 wherein the structural formula is
II:
15



Formula II

20 wherein R₁, R₄, R_{4a}, Y and R₃ are as described above.

7. A compound which is:
N-[5(S)-3-[4-[(1 α ,5 α ,6 β)-(6-cyanobicyclo[3.1.0]hexan-6-yl)]phenyl]-2-oxazolidin-5-
25 ylmethyl]acetamide,

1-[5(R)-3-[4-[(1 α ,5 α ,6 β)-(6-cyanobicyclo[3.1.0]hexan-6-yl)]phenyl]-2-oxooxazolidin-5-ylmethyl]-1,2,3-triazole,
 N-[5(S)-3-[4-[(1 α ,5 α ,6 β)-(3-t-butoxycarbonyl-6-cyano-3-azabicyclo[3.1.0]hexan-6-yl)]phenyl]-2-oxooxazolidin-5-ylmethyl]acetamide,
 5 N-[5(S)-3-[4-[(1 α ,5 α ,6 β)-(6-cyano-3-azabicyclo[3.1.0]hexan-6-yl)]phenyl]-2-oxooxazolidin-5-ylmethyl]acetamide ,
 1-[5(R)-3-[4-[(1 α ,5 α ,6 β)-(3-t-butoxycarbonyl-6-cyano-3-azabicyclo[3.1.0]hexan-6-yl)]phenyl]-2-oxooxazolidin-5-ylmethyl]-1,2,3-triazole,
 1-[5(R)-3-[4-[(1 α ,5 α ,6 β)-(6-cyano-3-azabicyclo[3.1.0]hexan-6-yl)]phenyl]-2-oxooxazolidin-5-ylmethyl]-1,2,3-triazole ,
 10 N-[5(S)-3-[4-[(1 α ,5 α ,6 β)-(3-acetoxyacetyl-6-cyano-3-azabicyclo[3.1.0]hexan-6-yl)]phenyl]-2-oxooxazolidin-5-ylmethyl]acetamide,
 N-[5(S)-3-[4-[(1 α ,5 α ,6 β)-(6-cyano-3-hydroxyacetyl-3-azabicyclo[3.1.0]hexan-6-yl)]phenyl]-2-oxooxazolidin-5-ylmethyl]acetamide,
 N-[5(S)-3-[4-[(1 α ,5 α ,6 β)-(6-cyano-3-methanesulfonyl-3-azabicyclo[3.1.0]hexan-6-yl)]phenyl]-2-oxooxazolidin-5-ylmethyl]acetamide,
 15 N-[5(S)-3-[4-[(1 α ,5 α ,6 β)-(6-cyano-3-methyl-3-azabicyclo[3.1.0]hexan-6-yl)]phenyl]-2-oxooxazolidin-5-ylmethyl]acetamide,
 N-[5(S)-3-[4-[(1 α ,5 α ,6 β)-(3,6-dicyano-3-azabicyclo[3.1.0]hexan-6-yl)]phenyl]-2-
 20 oxooxazolidin-5-ylmethyl]acetamide,
 N-[5(S)-3-[4-[(1 α ,5 α ,6 β)-(6-cyano-3-cyanomethyl-3-azabicyclo[3.1.0]hexan-6-yl)]phenyl]-2-oxooxazolidin-5-ylmethyl]acetamide,
 5(R)-3-[4-[(1 α ,5 α ,6 β)-(3-t-butoxycarbonyl-6-cyano-3-azabicyclo[3.1.0]hexan-6-yl)]phenyl]-5-[(isoxazol-3-yl)oxy]methyloxazolidin-2-one,
 25 5(R)-3-[4-[(1 α ,5 α ,6 β)-(6-cyano-3-azabicyclo[3.1.0]hexan-6-yl)]phenyl]-5-[(isoxazol-3-yl)oxy]methyloxazolidin-2-one ,
 5(R)-3-[4-[(1 α ,5 α ,6 β)-(3-t-butoxycarbonyl-6-cyano-3-azabicyclo[3.1.0]hexan-6-yl)]phenyl]-5-[N-(t-butoxycarbonyl)-N-(isoxazol-3-yl)]aminomethyloxazolidin-2-one,
 5(R)-3-[4-[(1 α ,5 α ,6 β)-(6-cyano-3-azabicyclo[3.1.0]hexan-6-yl)]phenyl]-5-[N-(isoxazol-3-
 30 yl)]aminomethyloxazolidin-2-one,
 N-[5(S)-3-[4-[(1 α ,5 α ,6 β)-[6-cyano-3-(5-cyanopyridin-2-yl)-3-azabicyclo[3.1.0]hexan-6-yl]]phenyl]-2-oxooxazolidin-5-ylmethyl]acetamide,
 N-[5(S)-3-[4-[(1 α ,5 α ,6 β)-[6-cyano-3-(pyridin-2-yl)-3-azabicyclo[3.1.0]hexan-6-yl]]phenyl]-2-oxooxazolidin-5-ylmethyl]acetamide,

N-[5(S)-3-[4-[(1 α ,5 α ,6 β)-[3-acetyl-6-cyano-3-azabicyclo[3.1.0]hexan-6-yl]]phenyl]-2-oxooxazolidin-5-ylmethyl]acetamide,
N-[5(S)-3-[4-[(1 α ,5 α ,6 β)-[6-cyano-3-(pyrimidin-2-yl)-3-azabicyclo[3.1.0]hexan-6-yl]]phenyl]-2-oxooxazolidin-5-ylmethyl]acetamide,
5 N-[5(S)-3-[4-[(1 α ,5 α ,6 β)-[6-cyano-3-(4-pyridylmethyl)-3-azabicyclo[3.1.0]hexan-6-yl]]phenyl]-2-oxooxazolidin-5-ylmethyl]acetamide,
N-[5(S)-3-[4-[(1 α ,5 α ,6 β)-[6-cyano-3-(N-cyano-1-iminoethyl)-3-azabicyclo[3.1.0]hexan-6-yl]]phenyl]-2-oxooxazolidin-5-ylmethyl]acetamide,
N-[5(S)-3-[4-[(1 α ,5 α ,6 β)-[6-cyano-3-methoxycarbonyl-3-azabicyclo[3.1.0]hexan-6-
10 yl]]phenyl]-2-oxooxazolidin-5-ylmethyl]acetamide,
N-[5(S)-3-[4-[(1 α ,5 α ,6 β)-[6-cyano-3-(N-cyano-S-methylthioiminomethyl)-3-azabicyclo[3.1.0]hexan-6-yl]]phenyl]-2-oxooxazolidin-5-ylmethyl]acetamide,
N-[5(S)-3-[4-[(1 α ,5 α ,6 β)-[6-cyano-3-(N-cyanocarboxamidyl)-3-azabicyclo[3.1.0]hexan-6-
15 yl]]phenyl]-2-oxooxazolidin-5-ylmethyl]acetamide,
N-[5(S)-3-[4-[(1 α ,5 α ,6 β)-[3-(N,N'-t-butoxycarbonylcarboxamidyl)-6-cyano-3-azabicyclo[3.1.0]hexan-6-yl]]phenyl]-2-oxooxazolidin-5-ylmethyl]acetamide,
N-[5(S)-3-[4-[(1 α ,5 α ,6 β)-[3-carboxamidyl-6-cyano-3-azabicyclo[3.1.0]hexan-6-yl]]phenyl]-2-oxooxazolidin-5-ylmethyl]acetamide,
N-[5(S)-3-[4-[(1 α ,5 α ,6 β)-[3-(N-t-Butoxycarbonylamino)acetyl-6-cyano-3-
20 azabicyclo[3.1.0]hexan-6-yl]]phenyl]-2-oxooxazolidin-5-ylmethyl]acetamide,
N-[5(S)-3-[4-[(1 α ,5 α ,6 β)-[3-aminoacetyl-6-cyano-3-azabicyclo[3.1.0]hexan-6-yl]]phenyl]-2-oxooxazolidin-5-ylmethyl]acetamide,
N-[5(S)-3-[4-[(1 α ,5 α ,6 β)-[6-cyano-3-methanesulfonylacetyl-3-azabicyclo[3.1.0]hexan-6-
25 yl]]phenyl]-2-oxooxazolidin-5-ylmethyl]acetamide,
N-[5(S)-3-[4-[(1 α ,5 α ,6 β)-[6-cyano-3-(dibenzylphosphoryloxy)acetyl-3-azabicyclo[3.1.0]hexan-6-yl]]phenyl]-2-oxooxazolidin-5-ylmethyl]acetamide,
N-[5(S)-3-[4-[(1 α ,5 α ,6 β)-[6-cyano-3-(phosphoryloxy)acetyl-3-azabicyclo[3.1.0]hexan-6-
30 yl]]phenyl]-2-oxooxazolidin-5-ylmethyl]acetamide,
or their enantiomer, diastereomer, or pharmaceutically acceptable salt, hydrate or prodrug thereof wherein.

8. A pharmaceutical composition comprised of a compound in accordance with claim 1 in combination with a pharmaceutically acceptable carrier

and optionally a in combination with a vitamin selected from the group consisting of vitamin B2, vitamin B6, vitamin B12 and folic acid.

9. A method of treating or preventing a bacterial infection in a
5 mammalian patient in need thereof, comprising administering to said patient an effective amount of a compound of claim 1.

10. A method of treating or preventing bacterial infection or an oxazolidinone-associated side effect by administering an effective amount of a compound of formula I of claim 1 and an effective amount of one or more of a
10 vitamin selected from the group consisting of vitamin B2, vitamin B6, vitamin B12 and folic acid to a patient in need thereof.

11. A method according to claim 16 for treating or preventing oxazolidinone-associated normocytic anemia, peripheral sensory neuropathy, sideroblastic anemia, peripheral sensory neuropathy, optic neuropathy, seizures, thrombocytopenia, cheilosis, hypo-regenerative anemia, megaloblastic anemia and seborrheic dermatitis by administering an effective amount of vitamin B2 to a patient in need thereof.